

wherein R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{10} , R^{11} , R^{12} and R^{13} are each independently hydrogen, hydroxy, halogen, carboxy, optionally substituted lower alkyl, optionally substituted lower alkoxy, optionally substituted lower alkenyl, optionally substituted lower alkenyloxy, optionally substituted lower alkylthio, optionally substituted lower alkoxy carbonyl, optionally substituted acyloxy, optionally substituted lower alkylsulfonyl, optionally substituted lower alkylsulfonyloxy, optionally substituted lower alkylsulfinyl, nitro, cyano, formyl, optionally substituted amino, optionally substituted carbamoyl, optionally substituted sulfamoyl or optionally substituted heterocyclyl,

X is $-O-$, $-CH_2-$, $-NR^{14}-$ wherein R^{14} is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or acetyl, or $-S(O)_p-$ wherein p is an integer of 0 to 2,

Y is optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted lower alkynyl, optionally substituted acyl, optionally substituted cycloalkyl, optionally substituted cycloalkenyl, optionally substituted aryl or optionally

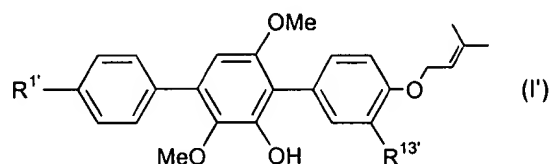
R¹ and R⁴, R¹ and R², R² and R³, R⁴ and R⁵, R⁶ and R⁷, R⁸ and R⁹, R¹⁰ and R¹¹, R¹² and R¹³, R¹¹ and -X-Y, or R¹³ and -X-Y taken together may form a 5- or 6-membered ring which may contain one or more of O, S or NR¹⁵ wherein R¹⁵ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted arylsulfonyl and which may optionally be substituted,

excluding compounds wherein one or more of R⁶, R⁷, R⁸ and R⁹ are halogen and the others are hydrogen, compounds wherein all of R⁶, R⁷, R⁸ and R⁹ are halogen and compounds wherein all of R²-R¹³ are hydrogen, halogen or cyano,

provided that R¹ is not hydrogen, fluorine, optionally substituted lower alkyl or optionally substituted lower alkoxy, all of R², R³, R⁴, R⁵ and R¹² are hydrogen, or R¹³ is not hydrogen or halogen when R⁶, R⁷, R⁸ and R⁹ are all simultaneously hydrogen,

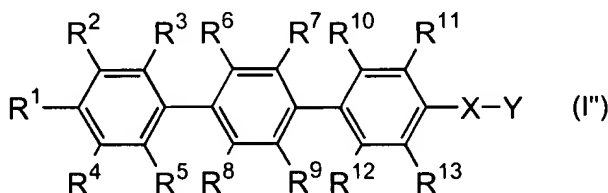
and further provided that R¹ is not methyl or acetyloxy, R¹³ is not hydrogen, optionally substituted lower alkoxy carbonyl or optionally

substituted carbamoyl, or -X-Y is not methoxy when at least one of R⁶, R⁷, R⁸ and R⁹ is a substituent other than hydrogen, and excluding a compound of the formula (I'):



wherein R^{1'} is hydrogen or hydroxy and R^{13'} is hydroxy or methoxy.

40. (Twice Amended) An immunosuppressive composition comprising a compound of the formula (I''):



wherein R¹, R², R³, R⁴, R⁵, R⁶, R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹² and R¹³ are each independently hydrogen, hydroxy, halogen, carboxy, optionally substituted lower alkyl optionally substituted, lower alkoxy, optionally substituted lower alkenyl, optionally substituted lower alkenyloxy, optionally substituted lower alkylthio, optionally substituted lower alkoxy carbonyl, optionally substituted acyloxy, optionally substituted lower alkylsulfonyl, optionally substituted lower alkylsulfonyloxy, optionally substituted lower alkylsulfinyl, nitro, cyano, formyl, optionally

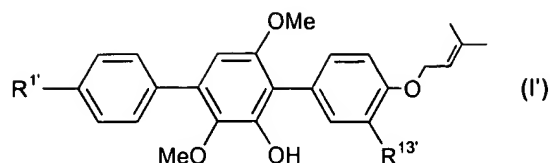
substituted amino, optionally substituted carbamoyl, optionally substituted sulfamoyl or optionally substituted heterocyclyl,

X is -O-, -CH₂-, -NR¹⁴- wherein R¹⁴ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or acetyl, or -S(O)_p- wherein p is an integer of 0 to 2,

Y is optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted lower alkynyl, optionally substituted acyl, optionally substituted cycloalkyl, optionally substituted cycloalkenyl, optionally substituted aryl or optionally substituted heterocyclyl, and Y may optionally be substituted lower alkoxy when X is -CH₂- and may optionally be substituted lower alkoxycarbonyl, optionally substituted lower alkylsulfonyl or optionally substituted arylsulfonyl when X is -O- or -NR¹⁴-,

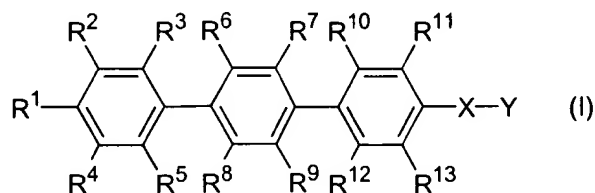
R¹ and R⁴, R¹ and R², R² and R³, R⁴ and R⁵, R⁶ and R⁷, R⁸ and R⁹, R¹⁰ and R¹¹, R¹² and R¹³, R¹¹ and -X-Y, or R¹³ and -X-Y taken together may form a 5- or 6-membered ring which may contain one or more of O, S or NR¹⁵ wherein R¹⁵ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or optionally substituted arylsulfonyl and which may optionally be substituted,

excluding a compound of the formula (I'):



F² wherein R^{1'} is hydrogen or hydroxy and R^{13'} is hydroxy or methoxy; or a pharmaceutically acceptable salt or hydrate or prodrug thereof, and a pharmaceutically acceptable excipient.

F³ 42. (Amended) A process for producing a compound of the formula (I):



wherein R¹, R², R³, R⁴, R⁵, R⁶, R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹² and R¹³ are each independently hydrogen, hydroxy, halogen, carboxy, optionally substituted lower alkyl, optionally substituted lower alkoxy, optionally substituted lower alkenyl, optionally substituted lower alkenyloxy, optionally substituted lower alkylthio, optionally substituted lower alkoxycarbonyl, optionally substituted acyloxy, optionally substituted lower alkylsulfonyl, optionally substituted lower alkylsulfonyloxy, optionally substituted lower alkylsulfinyl, nitro, cyano, formyl, optionally

substituted amino, optionally substituted carbamoyl, optionally substituted sulfamoyl or optionally substituted heterocyclyl,

X is -O-, -CH₂-, -NR¹⁴- wherein R¹⁴ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or acetyl, or -S(O)_p- wherein p is an integer of 0 to 2,

Y is optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted lower alkynyl, optionally substituted acyl, optionally substituted cycloalkyl, optionally substituted cycloalkenyl, optionally substituted aryl or optionally substituted heterocyclyl, and Y may optionally be substituted lower alkoxy when X is -CH₂- and may optionally be substituted lower alkoxycarbonyl, optionally substituted lower alkylsulfonyl or optionally substituted arylsulfonyl when X is -O- or -NR¹⁴-,

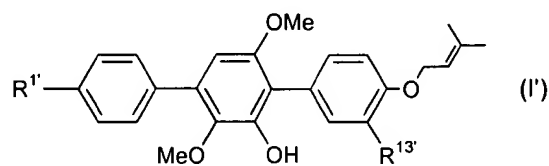
R¹ and R⁴, R¹ and R², R² and R³, R⁴ and R⁵, R⁶ and R⁷, R⁸ and R⁹, R¹⁰ and R¹¹, R¹² and R¹³, R¹¹ and -X-Y, or R¹³ and -X-Y taken together may form a 5- or 6-membered ring which may contain one or more of O, S or NR¹⁵ wherein R¹⁵ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted arylsulfonyl and which may optionally be substituted,

excluding compounds wherein one or more of R⁶, R⁷, R⁸ and R⁹ are halogen and the others are hydrogen, compounds wherein all of R⁶, R⁷, R⁸ and R⁹ are halogen and compounds wherein all of R²-R¹³ are hydrogen, halogen or cyano,

provided that R¹ is not hydrogen, fluorine, optionally substituted lower alkyl or optionally substituted lower alkoxy, all of R², R³, R⁴, R⁵ and R¹² are hydrogen, or R¹³ is not hydrogen or halogen when R⁶, R⁷, R⁸ and R⁹ are all simultaneously hydrogen,

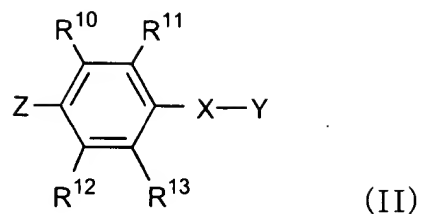
and further provided that R¹ is not methyl or acetyloxy, R¹³ is not hydrogen, optionally substituted lower alkoxy, carbonyl or optionally substituted carbamoyl, or -X-Y is not methoxy when at least one of R⁶, R⁷, R⁸ and R⁹ is a substituent other than hydrogen,

and excluding a compound of the formula (I'):

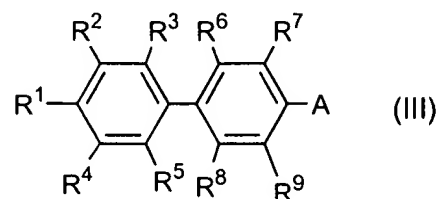


wherein R^{1'} is hydrogen or hydroxy and R^{13'} is hydroxy or methoxy;

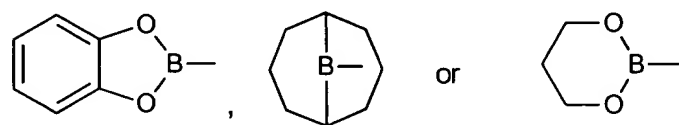
said process comprising reacting a compound of the formula (II):



3 with a compound of the formula (III):

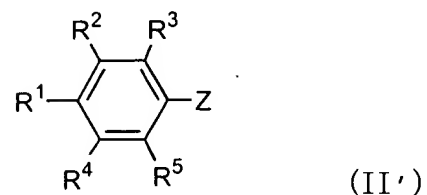


wherein, in the formulas (II) and (III), R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , R^7 , R^8 , R^9 , R^{10} , R^{11} , R^{12} and R^{13} are each described above; either of A and Z is dihydroxyborane, di(lower)alkoxyborane, di(lower)alkylborane,

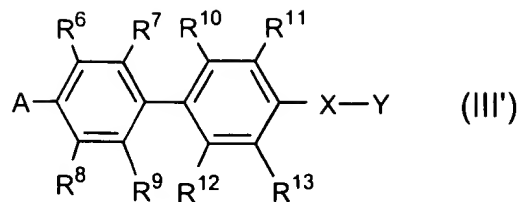


and the other is halogen or $-\text{OSO}_2(\text{C}_q\text{F}_{2q+1})-$ wherein q is an integer of 0 to 4,

or reacting a compound of the formula (II'):



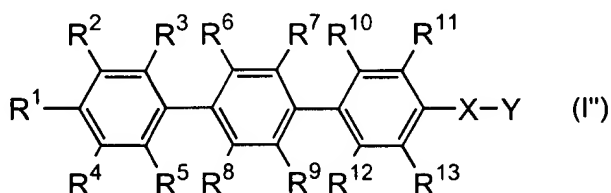
with a compound of the formula (III'):



wherein, in the formulas (II') and (III'), $R^1 - R^{13}$, X and Y are the same as defined above and A and Z are the same as defined in the above formulas (II) and (III).

F 4

56. (Amended) An immunosuppressive composition comprising a compound of the formula (I''):



wherein $R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^8, R^9, R^{10}, R^{11}, R^{12}$ and R^{13} are each independently hydrogen, hydroxy, carboxy, optionally substituted lower alkoxy, optionally substituted lower alkenyl, optionally substituted lower alkenyloxy, optionally substituted lower alkylthio, optionally substituted lower alkoxycarbonyl, optionally substituted acyloxy, optionally substituted lower alkylsulfonyl, optionally substituted lower alkylsulfonyloxy, optionally substituted lower alkylsulfinyl, nitro, formyl, optionally substituted amino, optionally substituted carbamoyl,

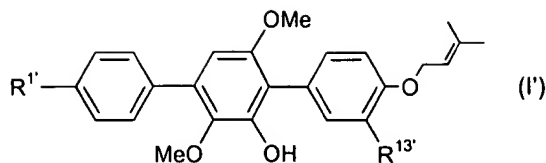
optionally substituted sulfamoyl or optionally substituted heterocyclyl,

X is -O-, -CH₂-, -NR¹⁴- wherein R¹⁴ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or acetyl, or -S(O)_p- wherein p is an integer of 0 to 2,

Y is optionally substituted lower alkyl, optionally substituted lower alkenyl, optionally substituted lower alkynyl, optionally substituted acyl, optionally substituted cycloalkyl, optionally substituted cycloalkenyl, optionally substituted aryl or optionally substituted heterocyclyl, and Y may optionally be substituted lower alkoxy when X is -CH₂- and may optionally be substituted lower alkoxycarbonyl, optionally substituted lower alkylsulfonyl or optionally substituted arylsulfonyl when X is -O- or -NR¹⁴-,

R¹ and R⁴, R¹ and R², R² and R³, R⁴ and R⁵, R⁶ and R⁷, R⁸ and R⁹, R¹⁰ and R¹¹, R¹² and R¹³, R¹¹ and -X-Y, or R¹³ and -X-Y taken together may form a 5- or 6-membered ring which may contain one or more of O, S or NR¹⁵ wherein R¹⁵ is hydrogen, optionally substituted lower alkyl, optionally substituted lower alkenyl or optionally substituted arylsulfonyl and which may optionally be substituted,

excluding a compound of the formula (I'):



wherein R^{1'} is hydrogen or hydroxy and R^{13'} is hydroxy or methoxy; or a pharmaceutically acceptable salt or hydrate or prodrug thereof, and a pharmaceutically acceptable excipient.

Please add the following new claims:

F 5

--57. (NEW) The compound or immunosuppressive composition as claimed in any one of claims 34, 40, 45, 46 and 49, wherein at least two of R⁶, R⁷, R⁸ and R⁹, are each independently selected from the group consisting of hydroxy, carboxy, optionally substituted lower alkyl, optionally substituted lower alkoxy, optionally substituted lower alkenyl, optionally substituted lower alkenyloxy, optionally substituted lower alkylthio, optionally substituted lower alkoxycarbonyl, optionally substituted acyloxy, optionally substituted lower alkylsulfonyl, optionally substituted lower alkylsulfonyloxy, optionally substituted lower alkylsulfinyl, nitro, cyano, formyl, optionally substituted amino, optionally substituted carbamoyl, optionally substituted sulfamoyl and optionally substituted heterocyclyl.

58. (NEW) The compound claimed in claim 57, wherein at least
⁵ one of R⁶, R⁷, R⁸ and R⁹ is a hydrogen, optionally substituted
lower alkoxy or optionally substituted lower alkyl.
